

RECEIVED PCTO 19 APR 2006  
EV66 449000USATTORNEY DOCKET NUMBER  
*#4*

NEN-22902/16

U.S. APPLICATION NO. (If known, see 37 CFR 1.5)  
10/539,273INTERNATIONAL APPLICATION NO. INTERNATIONAL FILING DATE  
PCT/US04/028238 30 August 2004PRIORITY DATE CLAIMED  
29 August 2003

## TITLE OF INVENTION

MASS SPECTROMETRY METHODS FOR SIMULTANEOUS DETECTION OF METABOLIC ENZYME ACTIVITY AND METABOLITE LEVELS

## APPLICANT(S) FOR DO/EO/US

CERDA, Blas

Applicant herewith submits to the United States Designated/Elected Office (DO/EO/US) the following items and other information:

1.  This is a **FIRST** submission of items concerning a submission under 35 U.S.C. 371.
2.  This is a **SECOND** or **SUBSEQUENT** submission of items concerning a submission under 35 U.S.C. 371.
3.  This is an express request to begin national examination procedures (35 U.S.C. 371(f)). The submission must include items (5), (6), (9) and (24) indicated below.
4.  The US has been elected (Article 31).
5.  A copy of the International Application as filed (35 U.S.C. 371 (c)(2))
  - a.  is attached hereto (required only if not communicated by the International Bureau).
  - b.  has been communicated by the International Bureau.
  - c.  is not required, as the application was filed in the United States Receiving Office (RO/US).
6.  An English language translation of the International Application as filed (35 U.S.C. 371(c)(2))
  - a.  is attached hereto.
  - b.  has been previously submitted under 35 U.S.C. 154(d)(4).
7.  Amendments to the claims of the International Application under PCT Article 19 (35 U.S.C. 371 (c)(3))
  - a.  are attached hereto (required only if not communicated by the International Bureau).
  - b.  have been communicated by the International Bureau.
  - c.  have not been made; however, the time limit for making such amendments has NOT expired.
  - d.  have not been made and will not be made.
8.  An English language translation of the amendments to the claims under PCT Article 19 (35 U.S.C. 371(c)(3)).
9.  An oath or declaration of the inventor(s) (35 U.S.C. 371 (c)(4)).
10.  An English language translation of the annexes to the International Preliminary Examination Report under PCT Article 36 (35 U.S.C. 371 (c)(5)).
11.  A copy of the International Preliminary Examination Report (PCT/IPEA/409).
12.  A copy of the International Search Report (PCT/ISA/210).

## Items 13 to 23 below concern document(s) or information included:

13.  An Information Disclosure Statement under 37 CFR 1.97 and 1.98.
14.  An assignment document for recording. A separate cover sheet in compliance with 37 CFR 3.28 and 3.31 is included.
15.  A **FIRST** preliminary amendment.
16.  A **SECOND** or **SUBSEQUENT** preliminary amendment.
17.  A substitute specification.
18.  A power of attorney and/or change of address letter.
19.  A computer-readable form of the sequence listing in accordance with PCT Rule 13ter.2 and 37 CFR 1.821 - 1.825.
20.  A second copy of the published International Application under 35 U.S.C. 154(d)(4).
21.  A second copy of the English language translation of the International Application under 35 U.S.C. 154(d)(4).
22.  Express Mail Label No. EV 669544900 US

Under the Paperwork Reduction Act of 1995, no persons are required to respond to a collection of information unless it displays a valid OMB control number.

U.S. APPLICATION NO (if known, see 37 CFR 1.5)	INTERNATIONAL APPLICATION NO.	ATTORNEY'S DOCKET NUMBER
10/539,273	PCT/US04/028238	NEN-22902/16

## 23. Other items or information:

Copy of Notification of Missing Requirements Under 35 U.S.C. 371 in the United States Designated/Elected Office (DO/EO/US)

Petition for Extension of Time Under 37 CFR 1.136(a) (Large Entity)

Check for \$580 (\$130 for surcharge fee and \$450 for 2 mo. extension fee)

The following fees have been submitted:				CALCULATIONS	PTO USE
24. <input type="checkbox"/> Basic national fee .....	\$300	\$	\$0.00		
25. <input type="checkbox"/> Examination fee (37 CFR 1.492(c)) If the written opinion prepared by ISA/US or the international preliminary examination report prepared by IPEA/US indicates all claims satisfy provisions of PCT Article 33(1)-(4). . . . .	\$0	\$	\$0.00		
All other situations. ....	\$200				
26. <input type="checkbox"/> Search fee (37 CFR 1.492(b)) If the written opinion of the ISA/US or the International preliminary examination report by IPEA/US indicates all claims satisfy provisions of PCT Article 33(1)-(4). . . . . Search fee (37 CFR 1.445(a)(2)) has been paid on the international application to the as an International Searching Authority. ....	\$0	\$	\$0.00		
International Search Report prepared by an ISA other than the US and provided to the previously communicated to the US by the IB. ....	\$100				
All other situations. ....	\$400				
<b>TOTAL OF 24, 25 and 26 =</b>	<b>\$500</b>	<b>\$</b>	<b>\$0.00</b>		
<input type="checkbox"/> Additional fee for specification and drawings filed in paper over 100 sheets (excluding sequence listing in compliance with 37 CFR 1.821(c) or (e) or computer program listing in an electronic medium) (37 CFR 1.492(j)). The fee is \$250 for each additional 50 sheets of paper or fraction thereof.					
Total Sheets	Extra Sheets	Number of each additional 50 or fraction thereof (round up to a whole)	RATE		
- 100 =	0 /50 =	0	x \$250.00	\$	\$0.00
Surcharge of \$130.00 for furnishing any of the search fee, examination fee, or the oath or declaration after the date of commencement of the national stage (37 CFR 1.492(h)).				\$	\$130.00
CLAIMS	NUMBER FILED	NUMBER EXTRA	RATE		
Total claims	- 20 =	0	x \$50.00	\$	\$0.00
Independent claims	- 3 =	0	x \$200.00	\$	\$0.00
MULTIPLE DEPENDENT CLAIMS (if applicable)	<input type="checkbox"/>	+ \$360.00		\$	\$0.00
<b>TOTAL OF ABOVE CALCULATIONS =</b>				\$	\$130.00
<input type="checkbox"/> Applicant claims small entity status. See 37 CFR 1.27. Fees above are reduced by 1/2.				\$	\$0.00
<b>SUBTOTAL =</b>				\$	\$130.00
Processing fee of \$130.00 for furnishing the English translation later than 30 months from the earliest claimed priority date (37 CFR 1.492(i)).				\$	\$0.00
<b>TOTAL NATIONAL FEE =</b>				\$	\$130.00
Fee for recording the enclosed assignment (37 CFR 1.21(h)). The assignment must be accompanied by an appropriate cover sheet (37 CFR 3.28, 3.31). \$40 per property +				\$	\$0.00
04/24/2006 LL ANDGRA 00000037 10539273	<b>TOTAL FEES ENCLOSED =</b>			\$	\$130.00
01 FC:1617	130.00 DP			Amount to be	\$
				Amount to be	\$

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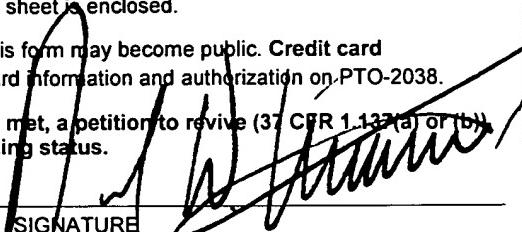
- a.  A check in the amount of \$ \$580.00 to cover the above fees is enclosed.
- b.  Please charge my Deposit Account No. \_\_\_\_\_ in the amount of \$ \_\_\_\_\_ to cover the above fees.  
A duplicate copy of this sheet is enclosed.
- c.  The Commissioner is hereby authorized to charge any additional fees which may be required, or credit any overpayment to Deposit Account No. 07-1180. A duplicate copy of this sheet is enclosed.
- d.  Fees are to be charged to a credit card. **WARNING:** Information on this form may become public. Credit card information should not be included on this form. Provide credit card information and authorization on PTO-2038.

NOTE: Where an appropriate time limit under 37 CFR 1.495 has not been met, a petition to revive (37 CFR 1.123(a) or (b)) must be filed and granted to restore the International Application to pending status.

SEND ALL CORRESPONDENCE TO:

Ronald W. Citkowski  
Gifford, Krass, Groh, Sprinkle,  
Anderson & Citkowski, P.C.  
P.O. Box 7021  
Troy, MI 48007-7021

SIGNATURE

  
Ronald W. Citkowski

NAME

31,005

REGISTRATION NUMBER

April 19, 2006

DATE



## UNITED STATES PATENT AND TRADEMARK OFFICE

UNITED STATES DEPARTMENT OF COMMERCE  
 United States Patent and Trademark Office  
 Address: COMMISSIONER FOR PATENTS  
 P.O. Box 1450  
 Alexandria, Virginia 22313-1450  
[www.uspto.gov](http://www.uspto.gov)

U.S. APPLICATION NUMBER NO.	FIRST NAMED APPLICANT	ATTY. DOCKET NO.
10/539,273	Blas Cerdá	Nen-22902/16
INTERNATIONAL APPLICATION NO.		
PCT/US04/28238		
I.A. FILING DATE	PRIORITY DATE	
08/30/2004	08/29/2003	

37742  
 GIFFORD, KRASS, GROH, SPRINKLE, ANDERSON  
 & CITKOWSKI, P.C.  
 P.O. BOX 7021  
 TROY, MI 48007-7021

CONFIRMATION NO. 8503

371 FORMALITIES LETTER



\*OC000000017681066\*

Date Mailed: 12/20/2005

**NOTIFICATION OF MISSING REQUIREMENTS UNDER 35 U.S.C. 371 IN THE UNITED STATES DESIGNATED/ELECTED OFFICE (DO/EO/US)**

The following items have been submitted by the applicant or the IB to the United States Patent and Trademark Office as a Designated / Elected Office (37 CFR 1.495).

- Copy of the International Application filed on 06/16/2005
- Preliminary Amendments filed on 06/16/2005
- Request for Immediate Examination filed on 06/16/2005
- U.S. Basic National Fees filed on 06/16/2005
- Priority Documents filed on 06/16/2005
- Power of Attorney filed on 06/16/2005

The following items **MUST** be furnished within the period set forth below in order to complete the requirements for acceptance under 35 U.S.C. 371:

- Oath or declaration of the inventors, in compliance with 37 CFR 1.497(a) and (b), identifying the application by the International application number and international filing date.

**ALL OF THE ITEMS SET FORTH ABOVE MUST BE SUBMITTED WITHIN TWO (2) MONTHS FROM THE DATE OF THIS NOTICE OR BY 32 MONTHS FROM THE PRIORITY DATE FOR THE APPLICATION, WHICHEVER IS LATER. FAILURE TO PROPERLY RESPOND WILL RESULT IN ABANDONMENT.**

The time period set above may be extended by filing a petition and fee for extension of time under the provisions of 37 CFR 1.136(a).

Applicant is reminded that any communications to the United States Patent and Trademark Office must be mailed to the address given in the heading and include the U.S. application no. shown above (37 CFR 1.5)

*A copy of this notice **MUST** be returned with the response.*

R E C E I V E  
 DEC 27 2005

GIFFORD, KRASS, GROH, SPRINKLE,  
 ANDERSON & CITKOWSKI, P.C.

SHELBY J VIGIL

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Telephone: (703) 308-9140 EXT 224

PART 1 - ATTORNEY/APPLICANT COPY

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U.S. APPLICATION NUMBER NO.	INTERNATIONAL APPLICATION NO.	ATTY. DOCKET NO.
10/539,273	PCT/US04/28238	Nen-22902/16

FORM PCT/DO/EO/905 (371 Formalities Notice)